$$\begin{array}{c|c} XH \cdot H_2N & H & H \\ \hline \\ O & \\ \hline \\ CO_2R^1 & \end{array}$$
 (I)

wherein R1 is para-nitrobenzy or allyl; and X is halo;

comprising the steps of

a) cyclizing a trimethylphosphinic compound of formula (IIIa)

$$R^2$$
 $C$ 
 $N$ 
 $P(CH_3)_3$ 
 $CO_2R^1$ 
(IIIa)

wherein

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R<sup>1</sup> is *para*-nitrobenzyl or allyl;

R<sup>2</sup> is selected from the group consisting of C<sub>1-6</sub>alkyl, C<sub>6-10</sub>aryl, C<sub>6-10</sub>arylC<sub>1-6</sub>alkyl and dithianyl;

in a solvent;

to form a compound of formula (II)

15 wherein

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R<sup>1</sup> is *para*-nitrobenzyl or allyl;

 $R^2$  is selected from the group consisting of  $C_{1-6}$ alkyl,  $C_{6-10}$ aryl,  $C_{6-10}$ aryl $C_{1-6}$ alkyl and dithianyl; and

- b) reacting said compound of formula (II) with an acid.
- 2. A process according to claim 1, wherein said solvent is selected from the group consisting of toluene, xylene, tetrahydrofuran, methylene chloride and acetonitrile.

  USERSIDOCS\LA21952\LPEXD\JCC6011.DOC/155958/PC10856.EXD

- 3. A process according to claim 1, wherein said acid is phosphorus pentachloride or phosphorus pentabromide; and wherein X is chloro or bromo.
- 4. A process according to claim 1, further comprising the step of preparing said compound of formula (IIIa), by reacting a compound of formula (IIIb)

$$R^2$$
 $N$ 
 $N$ 
 $X$ 
 $CO_2R^1$ 
(IIIb)

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wherein said R<sup>1</sup> is para-nitrobenzyl or allyl,

said  $R^2$  is selected from the group consisting of  $C_{1-6}$ alkyl,  $C_{6-10}$ aryl,  $C_{6-10}$ aryl $C_{1-6}$ alkyl and dithianyl; and

said X is halo;

with trimethylphosphine, in a solvent and in the presence of a base.

- 5. A process according to claim 4, wherein said solvent is tetrahydrofuran, acetonitrile or methylene chloride.
- 6. A process according to claim 4, wherein said base is selected from the group consisting of imidazole, 2,6-lutidine, pyridine, N-methylmorpholine and sodium bicarbonate.
- 7. A process according to claim 4, further comprising the step of preparing said compound of formula (IIIb), by reacting a compound of formula (IIIc)

$$\begin{array}{c|c}
R^2 & H & H & H \\
\hline
N & H & H & S \\
\hline
O & O & O \\
\hline
CO_2R^1 & (IIIc)
\end{array}$$

wherein said  $R^1$  is *para*-nitrobenzyl or allyl and said  $R^2$  is selected from the group consisting of  $C_{1-6}$ alkyl,  $C_{6-10}$ aryl,  $C_{6-10}$ aryl $C_{1-6}$ alkyl and dithianyl; with a halogenating agent, in a solvent and in the presence of a base.

- 8. A process according to claim 7, wherein said halogenating agent is thionyl chloride, thionyl bromide, phosphorus trichloride or phosphorus tribromide; and said halo is chloro or bromo.
- 9. A process according to claim 7, wherein said base is selected from the group consisting of pyridine, 2,6-lutidine, N-methylmorpholine and imidazole.
- 10. A process according to claim 7, further comprising the step of preparing said compound of formula (IIIc), by reacting a compound of formula (V)

$$R^{2} \xrightarrow{C} \xrightarrow{N} \xrightarrow{H} \xrightarrow{H} \xrightarrow{H} \xrightarrow{H} SH$$

$$CO_{2}R^{1} \qquad (V)$$

wherein said  $R^1$  is *para*-nitrobenzyl or allyl and said  $R^2$  is selected from the group consisting of  $C_{1-6}$ alkyl,  $C_{6-10}$ aryl $C_{1-6}$ alkyl and dithianyl;

with a compound of formula (IV)

$$Y-CH_2$$
  $O$   $(IV)$ 

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wherein Y is a leaving group selected from the group consisting of bromo, chloro, fluoro, iodo and tosylate; in a solvent.

- 11. A process according to claim 10, wherein said Y is bromo or chloro.
- 12. A process according to claim 10 wherein said solvent is alcohol selected from the group consisting of methanol, ethanol and propanol; methylene chloride; acetone; dimethylformamide or mixtures thereof.
- 13. A process according to claim 10, further comprising the step of preparing said compound of formula (V) by reacting a compound of formula (VIa)

$$O \xrightarrow{R^2} O H CO_2 R^1$$
 (VIa)

- wherein  $R^1$  is *para*-nitrobenzyl or allyl and wherein  $R^2$  is selected from the group consisting of  $C_{1-6}$ alkyl,  $C_{6-10}$ aryl,  $C_{6-10}$ aryl $C_{1-6}$ alkyl and dithianyl; with an acid in a solvent.
  - 14. A process according to claim 13 wherein said acid is *para*-toluene sulfonic acid or methane sulfonic acid.
- 15. A process according to claim 13 wherein said solvent is methylene chloride,20 tetrahydrofuran, acetone or mixtures thereof.
  - 16. A process according to claim 13 further comprising the step of preparing said compound of formula (VIa) by:

reacting a compound of formula (VIb)

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wherein

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R<sup>1</sup> is para-nitrobenzyl or allyl;

 $R^2$  is selected from the group consisting of  $C_{1-6}$ alkyl,  $C_{6-10}$ aryl,  $C_{6-10}$ aryl $C_{1-6}$ alkyl and dithianyl;

with a reducing agent selected from the group consisting of sodium borohydride, sodium cyanoborohydride, borane and sodium triacetoxy borohydride; in a solvent.

- 17. A process according to claim 16 wherein said reducing agent is sodium triacetoxy borohydride.
- 10 18. A process according to claim 16 wherein said solvent is acetic acid, methylene chloride, tetrahydrofuran, isopropanol or mixtures thereof.
  - 19. A process according to claim 13 further comprising the step of preparing said compound of formula (VIa) by reacting a compound of formula (XI)

wherein R<sup>2</sup> is selected from the group consisting of C<sub>1-6</sub>alkyl, C<sub>6-10</sub>aryl, C<sub>6-10</sub>arylC<sub>1-6</sub>alkyl and dithianyl;

with a compound of formula (X)

$$OH$$
 $OR^1$ 
 $O$ 
 $O$ 
 $O$ 
 $O$ 

wherein R<sup>1</sup> is *para*-nitrobenzyl or allyl; in a solvent; in the presence of a base.

20. A process according to claim 16 further comprising the step of preparing said compound of formula (VIb) comprising reacting a compound of formula (VIII)

wherein

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 $R^2$  is selected from the group consisting of  $C_{1-6}$ alkyl,  $C_{6-10}$ aryl,  $C_{6-10}$ aryl $C_{1-6}$ alkyl and dithianyl;

 $L_2$  is a leaving group selected from the group consisting of halo, azide and  $C_{1-6}$ alkoxy; with a compound of formula (VII)

$$R^{1}$$
—OH (VII)

wherein R<sup>1</sup> is *para*-nitrobenzyl or allyl, in a solvent, in the presence of a base;

further comprising the step of preparing said compound of formula (VIII) by reacting a compound of formula (XI)

wherein  $R^2$  is selected from the group consisting of  $C_{1-6}$ alkyl,  $C_{6-10}$ aryl,  $C_{6-10}$ aryl  $C_{1-6}$ alkyl and dithianyl; with a compound of formula (IX)

- wherein each of said  $L_1$  and  $L_2$  is a leaving group selected from the group consisting of halo, azide and  $C_{1-6}$  alkoxy; in a solvent, optionally in the presence of a base.
  - 21. A process according to claim 16 further comprising the step of preparing said compound of formula (VIb) comprising reacting a compound of formula (VIc)

$$\begin{array}{c|c}
R^2 \\
S \\
CO_2R^1
\end{array}$$
(VIc)

wherein

R<sup>1</sup> is para-nitrobenzyl or allyl;

 $R^2$  is selected from the group consisting of  $C_{1-6}$ alkyl,  $C_{6-10}$ aryl,  $C_{6-10}$ aryl $C_{1-6}$ alkyl and dithianyl;

R<sup>3</sup> is hydrogen or C<sub>1-6</sub>alkyl; and

R<sup>4</sup> is hydrogen or C<sub>1-6</sub>alkyl; with ozone, in a solvent.

22. A process according to claim 16 further comprising the step of preparing said compound of formula (VIb) comprising reacting a compound of formula (XI)

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wherein  $R^2$  is selected from the group consisting of  $C_{1-6}$ alkyl,  $C_{6-10}$ aryl,  $C_{6-10}$ aryl,  $C_{1-6}$ alkyl, and dithianyl; with a compound of formula (XII)

$$\bigcup_{L_3}^{O} OR^1$$

wherein

said L<sub>3</sub> is halo;

R<sup>1</sup> is para-nitrobenzyl or allyl;

in a solvent, in the presence of a base.

- 23. A process according to claim 20, wherein each of  $L_1$  and  $L_2$ , wherever each of them occurs, is halo selected from the group consisting of bromo or chloro.
  - 24. A process according to claim 21 wherein R<sup>3</sup> is methyl and R<sup>4</sup> is methyl.
- 25. A process according to claim 7 wherein said solvent, wherever it occurs, is methylene chloride, tetrahydrofuran or mixtures thereof.

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- 26. A process according to claim 21 wherein said solvent is methylene chloride, tetrahydrofuran, isopropanol or mixtures thereof.
- 27. A process according to claim 19 wherein said base is selected from the group consisting of diisopropylamine, triethylamine, pyridine and 2,6-lutidine.
- 28. A process according to claim 1, wherein each of said R<sup>1</sup>, wherever it occurs, is *para*-nitrobenzyl.
  - 29. A process according to claim 1, wherein each of said R<sup>1</sup>, wherever it occurs, is allyl.
- 30. A process according to claim 1, wherein each of said  $R^2$ , wherever it occurs, 10 is  $C_{6-10}$ aryl $C_{1-6}$ alkyl.
  - 31. A process according to claim 1, wherein each of said R<sup>2</sup>, wherever it occurs, is benzyl.
    - 32. A compound of formula (I)

$$XH \cdot H_2N \stackrel{H}{\longrightarrow} H$$

$$CO_2R^1$$
(I)

- wherein R<sup>1</sup> is *para*-nitrobenzyl or allyl; and X is halo.
  - 33. A compound of formula (II)

$$R^2$$
  $C$   $HN$   $H$   $S$   $CO_2R^1$  (II)

wherein  $R^1$  is para-nitrobenzyl or allyl; and  $R^2$  is  $(C_6-C_{10})$ aryl $(C_{1-6})$ alkyl.

34. A compound of formula (III)

$$R^2$$
 $CO_2R^1$ 
(III)

wherein R1 is para-nitrobenzyl or allyl;

 $R^2$  is  $(C_6-C_{10})$ aryl $(C_{1-6})$ alkyl;

K is hydroxy, halo or -P-(CH<sub>3</sub>)<sub>3</sub>;

wherein the C-K bond is a single bond when K is hydroxy or halo; and a double bond when K is -P- $(CH_3)_3$ ; and

wherein said compound of formula (III) is selected from the group consisting of compound of formulae (IIIa), (IIIb) and (IIIc):

$$R^{2} \xrightarrow{HN} \xrightarrow{H} \xrightarrow{H} \xrightarrow{H} \xrightarrow{S} \xrightarrow{O} \xrightarrow{O} \xrightarrow{N} \xrightarrow{R^{2}} \xrightarrow{CO_{2}R^{1}} \xrightarrow{\text{(IIIb)}} \text{and}$$

10 35. A compound of formula (V)

wherein  $R^1$  is para-nitrobenzyl or allyl; and  $R^2$  is  $(C_6 - C_{10})$ aryl $(C_{1-6})$ alkyl.

36. A compound of formula (VI)

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wherein R<sup>1</sup> is para-nitrobenzyl or allyl;

 $R^2$  is  $(C_6-C_{10})$ aryl $(C_{1-6})$ alkyl;

T is hydroxy or >O;

wherein the C-T bond is a single bond when T is hydroxy; and a double bond when T is >O; and

wherein said compound of formula (VI) is selected from the group consisting of compound of formulae (VIa) and (VIb):

$$\bigcap_{O \to O \to O \to O}^{\mathbb{R}^2} \bigcap_{O \to O \to O}^{\mathbb{R}^2} \bigcap_{O \to O \to O}^{\mathbb{R}^2} \bigcap_{O \to O}^{\mathbb{R$$

- 37. A compound according to claim 32, wherein said R<sup>1</sup> is *para*-nitrobenzyl.
  - 38. A compound according to claim 32, wherein said R<sup>1</sup> is allyl.
  - 39. A compound according to claim 32, wherein said R<sup>2</sup> is benzyl.